#1

10/533,683

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PASSWORD:

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Welcome to STN International
NEWS
      1
                 Web Page for STN Seminar Schedule - N. America
         OCT 02
                 CA/CAplus enhanced with pre-1907 records from Chemisches
NEWS
                 Zentralblatt
NEWS
         OCT 19
                 BEILSTEIN updated with new compounds
NEWS
         NOV 15
                 Derwent Indian patent publication number format enhanced
NEWS
         NOV 19
                 WPIX enhanced with XML display format
NEWS
        NOV 30
                 ICSD reloaded with enhancements
NEWS 7
         DEC 04
                 LINPADOCDB now available on STN
NEWS 8
         DEC 14
                 BEILSTEIN pricing structure to change
NEWS 9
         DEC 17
                 USPATOLD added to additional database clusters
NEWS 10
         DEC 17
                 IMSDRUGCONF removed from database clusters and STN
         DEC 17
                 DGENE now includes more than 10 million sequences
NEWS 11
         DEC 17
                 TOXCENTER enhanced with 2008 MeSH vocabulary in
NEWS 12
                 MEDLINE segment
NEWS 13
         DEC 17
                 MEDLINE and LMEDLINE updated with 2008 MeSH vocabulary
NEWS 14
         DEC 17
                 CA/CAplus enhanced with new custom IPC display formats
NEWS 15
         DEC 17
                 STN Viewer enhanced with full-text patent content
                 from USPATOLD
NEWS 16
         JAN 02
                 STN pricing information for 2008 now available
NEWS 17
         JAN 16
                 CAS patent coverage enhanced to include exemplified
                 prophetic substances
NEWS 18
         JAN 28
                 USPATFULL, USPAT2, and USPATOLD enhanced with new
                 custom IPC display formats
NEWS 19
         JAN 28
                 MARPAT searching enhanced
NEWS 20
         JAN 28
                 USGENE now provides USPTO sequence data within 3 days
                 of publication
NEWS 21
         JAN 28
                 TOXCENTER enhanced with reloaded MEDLINE segment
NEWS 22
         JAN 28
                 MEDLINE and LMEDLINE reloaded with enhancements
NEWS 23
        FEB 08
                 STN Express, Version 8.3, now available
NEWS 24
         FEB 20
                 PCI now available as a replacement to DPCI
                 IFIREF reloaded with enhancements
NEWS 25
         FEB 25
NEWS 26 FEB 25
                 IMSPRODUCT reloaded with enhancements
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NEWS EXPRESS FEBRUARY 08 CURRENT WINDOWS VERSION IS V8.3,
AND CURRENT DISCOVER FILE IS DATED 20 FEBRUARY 2008

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NEWS LOGIN Welcome Banner and News Items
NEWS IPC8 For general information regarding STN implementation of IPC 8

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FILE 'HOME' ENTERED AT 12:34:22 ON 25 FEB 2008

=> FILE REG COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

FULL ESTIMATED COST

FILE 'REGISTRY' ENTERED AT 12:34:44 ON 25 FEB 2008
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STRUCTURE FILE UPDATES: 24 FEB 2008 HIGHEST RN 1005323-41-0 DICTIONARY FILE UPDATES: 24 FEB 2008 HIGHEST RN 1005323-41-0

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TSCA INFORMATION NOW CURRENT THROUGH January 9, 2008.

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http://www.cas.org/support/stngen/stndoc/properties.html

=> Uploading C:\Program Files\Stnexp\Queries\LC-52.str

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chain nodes :
13 14 15 16 17 18 19 20 21
                               22
                                  23
                                      24 25 26 27 28 29 30 31
34 35 36
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12
chain bonds :
1-19 2-31 3-17 4-29 5-30 6-13 7-35 8-34 9-33 10-32 11-19 12-36 13-14
14-15 14-16 17-18 19-20 20-21 21-22 22-23 22-26 23-24 23-25 26-27 26-28
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12
exact/norm bonds :
6-13 13-14 14-15 14-16 17-18 22-23 22-26
exact bonds :
1-19 2-31 3-17 4-29 5-30 7-35 8-34 9-33 10-32 11-19 12-36 19-20 20-21
21-22 23-24 23-25 26-27 26-28
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12
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# G1:Cb,Cy,Ak

Match level:
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS
19:CLASS 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS
27:CLASS 28:CLASS 29:CLASS 30:CLASS 31:CLASS 32:CLASS 33:CLASS 34:CLASS
35:CLASS 36:CLASS

#### STRUCTURE UPLOADED

=> D L1

T.1

L1 HAS NO ANSWERS

1.1

STR

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Structure attributes must be viewed using STN Express query preparation.

=> S L1 FULL

FULL SEARCH INITIATED 12:35:32 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 84 TO ITERATE

100.0% PROCESSED

84 ITERATIONS

19 ANSWERS

SEARCH TIME: 00.00.01

L2

19 SEA SSS FUL L1

=> FILE CAPLUS

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

178.82

179.03

FILE 'CAPLUS' ENTERED AT 12:35:42 ON 25 FEB 2008
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=> S L2

L3 31 L2

=> S L2 AND DEVICE

31 L2

901259 DEVICE

L4 1 L2 AND DEVICE

=> S L3 AND DEVICE

901259 DEVICE

L5 1 L3 AND DEVICE

=> S L3 AND TRANSDERMAL

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=> S L3 AND DELIVERY
289142 DELIVERY
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L7 14 L3 AND DELIVERY

=> D L7 IBIB ABS HITSTR 1-14

L7 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2008:70709 CAPLUS

DOCUMENT NUMBER: 148:152045

TITLE: Pharmaceutical preparation for oral administration

with controlled active ingredient release in the small

intestine and methods for its production

INVENTOR(S): Jung, Gerd; Schaupp, Albert

PATENT ASSIGNEE(S): Dr. R. Pfleger Chemische Fabrik GmbH, Germany

SOURCE: PCT Int. Appl., 41pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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PATENT NO.
                   KIND
                          DATE
                                    APPLICATION NO.
                                                            DATE
                                    ______
                         _____
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                                    WO 2007-EP5970
WO 2008006506
                         20080117
                   A1
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       GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG,
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       BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW,
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       BY, KG, KZ, MD, RU, TJ, TM
EP 1880718
                   A1
                         20080123
                                     EP 2006-14244
                                                            20060710
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       IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL,
       BA, HR, MK, YU
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PRIORITY APPLN. INFO.: EP 2006-14244 A 20060710 · A pharmaceutical preparation for oral administration with controlled active ingredient release in the small intestine, on the basis of active ingredient carriers provided with at least one active ingredient which are provided with an inner layer for controlling the active ingredient release and a covering layer, arranged thereon, that is resistant to gastric juices, and is characterized in that the inner layer is constructed from at least two diffusion layers whose permeability for the diffusing active ingredient decreases from the inside to the outside, and a method for its production are described. Thus (1R, 3R, 5S)-3-[(Hydroxydiphenylacetyl)oxy]spiro [8-azoniabicyclo[3.2.1]octane-8,1'-pyrrolidinium] chloride-containing pharmaceutical formulations were prepared Pellets contained mg/dose: drug 45.000; neutral pellets 100.000; hypromellose 4.500; Macrogol 6000 0.450; total 154.450. The first diffusion layer was applied onto the above pellets, mg/dose: drug pellet 154.450; Kollicoat SR 30D 9.000; Kollicoat IR 1.800; propyleneglycol 0.900; talc 0.360; total 166.510. The second diffusion layer was applied onto the above coated pellets, mg/dose: drug pellet 166.510; Kollicoat SR 30D 9.000; Kollicoat IR 1.800; propyleneglycol 0.900; talc 0.360; total 177.175. The gastric juice resistant layer was applied onto the above coated pellets, mg/dose: drug pellet (containing 45 mg drug) 177.175, Kollicoat MAE30DP 28.000; talc 12.600; propylene glycol 4.200; Tylopur C30G1 0.720; total 222.695.

IT 286930-02-7

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (pharmaceutical preparation for oral administration with controlled active ingredient release in small intestine and methods for its production)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2007:1454781 CAPLUS

DOCUMENT NUMBER:

148:78876

TITLE:

Cyclopentylpyrrolidinone derivatives and their

preparation and use in combination therapy for the treatment of urinary frequency, urinary urgency and urinary incontinence

INVENTOR(S): Go

Gottesdiener, Keith M.; Green, Stuart A.; Macintyre,

Euan

PATENT ASSIGNEE(S):

Merck & Co., Inc., USA

SOURCE:

PCT Int. Appl., 86pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

GΙ

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| LAICMI | INFORMATION: |
|--------|--------------|
|        |              |
|        |              |

| PAT      | ENT                 | NO.      |      |     | KIN | D . | DATE |      |     | APPL | ICAT | ION : | NO.     |     | D    | ATE  | ,               |
|----------|---------------------|----------|------|-----|-----|-----|------|------|-----|------|------|-------|---------|-----|------|------|-----------------|
| WO.      | - <b></b> -<br>2007 | <br>1462 | 24   |     | A2  | -   | 2007 | 1221 | 1   | WO 2 | 007- | US13  | <br>683 |     | 2    | 0070 | <b>-</b><br>607 |
|          | W:                  | ΑE,      | AG,  |     |     |     | AU,  |      |     |      |      |       |         |     |      |      |                 |
|          |                     |          |      |     |     |     | CZ,  |      |     |      |      |       |         |     |      |      |                 |
|          |                     |          |      |     |     |     | GT,  |      |     |      |      |       |         |     |      |      |                 |
|          |                     |          |      |     |     |     | LA,  |      |     |      |      |       |         |     |      |      |                 |
|          |                     |          |      |     |     |     | MY,  |      |     |      |      |       |         |     |      |      |                 |
|          |                     | PT,      | RO,  | RS, | RU, | SC, | SD,  | SE,  | SG, | SK,  | SL,  | SM,   | sv,     | SY, | ТJ,  | TM,  | TN,             |
|          |                     |          |      |     |     |     | US,  |      |     |      |      |       |         |     |      |      | •               |
|          | RW:                 | ΑT,      | BE,  | BG, | CH, | CY, | CZ,  | DE,  | DK, | EE,  | ES,  | FΙ,   | FR,     | GB, | GR,  | HU,  | IE,             |
|          |                     | IS,      | ΙT,  | LT, | LU, | LV, | MC,  | MT,  | NL, | PL,  | PT,  | RO,   | SE,     | SI, | SK,  | TR,  | BF,             |
|          |                     | ВJ,      | CF,  | CG, | CI, | CM, | GΑ,  | GN,  | GQ, | GW,  | ML,  | MR,   | NE,     | SN, | TD,  | TG,  | BW,             |
|          |                     | GH,      | GM,  | KE, | LS, | MW, | ΜZ,  | NA,  | SD, | SL,  | SZ,  | ΤZ,   | UG,     | ZM, | ZW,  | AM,  | AZ,             |
|          |                     | BY,      | KG,  | ΚZ, | MD, | RU, | ТJ,  | TM   |     |      |      |       |         |     |      |      |                 |
| PRIORITY | APP:                | LN.      | INFO | .:  |     |     |      |      | . 1 | US 2 | 006- | 8127  | 43P     | 1   | P 20 | 0060 | 612             |

This invention concerns compns. for the treatment of urinary frequency, urinary urgency and urinary incontinence comprising a selected antagonist of the NK-1 receptor or a pharmaceutically acceptable salt thereof and an anti-muscarinic agent or a pharmaceutically acceptable salt thereof. This invention concerns combination therapy for urinary frequency, urinary urgency and urinary incontinence wherein one of the active agents is a selected antagonist of the NK-1 receptor or a pharmaceutically acceptable salt thereof and another is an anti-muscarinic agent or a pharmaceutically acceptable salt thereof. Example compound I was prepared by a multistep procedure (procedure given). All the invention compds. were evaluated for their NK-1 receptor antagonistic activity.

IT 286930-02-7, Fesoterodine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of cyclopentylpyrrolidinone derivs. as anti-muscarinic agents and NK-1 receptor antagonists in combination therapy of urinary frequency, urinary urgency and urinary incontinence)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L7 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN ACCESSION NUMBER: 2007:1425394 CAPLUS

DOCUMENT NUMBER:

148:45893

TITLE:

Treatment of excess sebum production

INVENTOR(S):

Roach, Alan George; Goldsmith, Paul

PATENT ASSIGNEE(S): SOURCE:

Daniolabs Ltd., UK PCT Int. Appl., 12pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT | TENT | ENT NO. |        |     | KIN | D    | DATE |      | j   | APPL | ICAT | ION  | NO.    |     | D.  | ATE  |         |
|-----|------|---------|--------|-----|-----|------|------|------|-----|------|------|------|--------|-----|-----|------|---------|
| WO  | 2007 | 1415    | <br>30 |     | A2  | -    | 2007 | 1213 | 1   | WO 2 | 007- | GB20 | <br>98 |     | 2   | 0070 | <br>607 |
|     | W:   | ΑE,     | AG,    | AL, | AM, | ΑT,  | ĂU,  | ΑZ,  | BA, | BB,  | ·BG, | BH,  | BR,    | BW, | BY, | ΒZ,  | CA,     |
|     |      | CH,     | CN,    | CO, | CR, | CU,  | CZ,  | DE,  | DK, | DM,  | DO,  | DZ,  | EC,    | EE, | EG, | ES,  | FI,     |
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|     |      | ΚM,     | KN,    | ΚP, | KR, | ΚZ,  | LA,  | LC,  | LK, | LR,  | LS,  | LT,  | LU,    | LY, | MA, | MD,  | MG,     |
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|     |      | TT,     | TZ,    | UA, | UG, | US,  | ŲΖ,  | VC,  | VN, | ZA,  | ZM,  | ZW   |        |     |     |      |         |
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|     |      | BY,     | KG,    | KΖ, | MD, | RU,  | ТJ,  | TM   |     |      |      |      |        |     |     |      |         |

PRIORITY APPLN. INFO.:

GB 2006-11240 A

A 20060607

AB A muscarinic receptor antagonist is useful for the treatment or prevention of a condition associated with excess sebum production or excretion.

Muscarinic

receptor antagonist oxybutynin dose-dependently reduced sebum production in healthy human volunteers.

IT 286930-02-7, Fesoterodine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(muscarinic receptor antagonist for treatment of excess sebum production)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L7 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2007:1420174 CAPLUS

DOCUMENT NUMBER:

148:62011

TITLE:

Stabilized pharmaceutical compositions comprising

fesoterodine

INVENTOR(S):

Arth, Christoph; Mika, Hans-Juergen; Komenda, Michael;

Lindner, Hans; Bicane, Fatima; Paulus, Kerstin;

Irngartiner, Meike

PATENT ASSIGNEE(S):

Schwarz Pharma AG, Germany

SOURCE:

PCT Int. Appl., 74pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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APPLICATION NO.
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     WO 2007141298·
                               20071213
                                          WO 2007-EP55582
                         A1
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     EP 1864651
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PRIORITY APPLN. INFO.:
                                           EP 2006-11941
                                                               A 20060609
                                           EP 2006-11942
                                                               A 20060609
                                           EP 2006-11943
                                                               A 20060609
AΒ
     The present application relates to a pharmaceutical composition comprising
     fesoterodine or a pharmaceutically acceptable salt or solvate thereof and
     a stabilizer selected from the group consisting of xylitol, sorbitol,
     cellulose 41.5, hypromellose (e.g. Methocel K100M) 70.0, hypromellose
```

polydextrose, isomalt and dextrose. A tablet contained fesoterodine hydrogen fumarate 4.0, xylitol 76.0, lactose monohydrate 43.0, microcryst. (e.g. Methocel K4M) 70.0, glycerol dibehenate 8.0, talc 7.5, and purified water q.s.

IT 286930-02-7, Fesoterodine 286930-03-8

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(stabilized pharmaceutical compns. comprising fesoterodine)

286930-02-7 CAPLUS RN

Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-CN phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 286930-03-8 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester, (2E)-2-butenedioate (1:1) (CA INDEX NAME)

CM 1

CRN 286930-02-7 CMF C26 H37 N O3

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-17-8 CMF C4 H4 O4

Double bond geometry as shown.

REFERENCE COUNT:

4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2007:940100 CAPLUS

DOCUMENT NUMBER:

147:269265

TITLE:

Combination of an  $\alpha 2$ -receptor agonist (such as clonidine) and an antimuscarinic agent (such as oxybutynin) for the treatment of sialorrhea

INVENTOR(S): Roach, Alan George; Goldsmith, Paul

PATENT ASSIGNEE(S):

Daniolabs Ltd., UK PCT Int. Appl., 16pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

SOURCE:

NT: 1

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT      | ENT 1 | NO.  |      |     | KIN | D 1 | DATE |      | Ž   | APPL | ICAT: | I NOI | . 01     |      | Di  | ATE   |     |
|----------|-------|------|------|-----|-----|-----|------|------|-----|------|-------|-------|----------|------|-----|-------|-----|
| WO :     | 2007  | 0938 | 24   |     | A1  | - : | 2007 | 0823 |     | WO 2 | 007-0 | GB500 | <b>-</b> |      | 2   | 0070: | 212 |
|          | W:    | ΑE,  | AG,  | AL, | AM, | AT, | ΑU,  | AZ,  | BA, | BB,  | BG,   | BR,   | BW,      | BY,  | ΒZ, | CA,   | CH, |
|          |       |      |      |     |     |     |      |      |     |      |       |       |          | ES,  |     |       |     |
|          |       |      |      |     |     |     |      |      |     |      |       |       |          | KΕ,  |     |       |     |
|          |       |      |      |     |     |     |      |      |     |      |       |       |          | MA,  |     |       |     |
|          |       | MN,  | MW,  | MX, | MY, | MZ, | NA,  | NG,  | NΙ, | NO,  | ΝZ,   | OM,   | PG,      | PH,  | PL, | PT,   | RO, |
|          |       | RS,  | RU,  | SC, | SD, | SE, | SG,  | SK,  | SL, | SM,  | SV,   | SY,   | ТJ,      | ·ΤΜ, | TN, | TR,   | TT, |
|          |       | TZ,  | UA,  | UG, | US, | UZ, | VC,  | VN,  | ZA, | ZM,  | ZW    |       |          |      |     |       |     |
|          | RW:   |      |      |     |     |     |      |      |     |      |       |       |          | GB,  |     |       |     |
|          |       | IS,  | IT,  | LT, | LU, | LV, | MC,  | NL,  | PL, | PT,  | RO,   | SE,   | SI,      | SK,  | TR, | BF,   | ВJ, |
|          |       | CF,  | CG,  | CI, | CM, | GΑ, | GN,  | GQ,  | GW, | ML,  | MR,   | ΝE,   | SN,      | TD,  | TG, | B₩,   | GH, |
|          |       | GM,  | ΚE,  | LS, | MW, | MZ, | NA,  | SD,  | SL, | SZ,  | TΖ,   | UG,   | ZM,      | ZW,  | AM, | ΑZ,   | BY, |
|          |       | KG,  | ΚZ,  | MD, | RU, | ТJ, | TM   |      |     |      |       |       |          |      |     |       |     |
| PRIORITY | APP   | LN.  | INFO | .:  |     |     |      |      | (   | GB 2 | 006-  | 2855  |          | 1    | A 2 | 0060  | 213 |
|          |       |      |      |     |     |     |      |      | 1   | GB 2 | 006-  | 2857  |          |      | A 2 | 0060  | 213 |

AB An  $\alpha 2$ -adrenoreceptor agonist (e.g. clonidine, brimonidine, monoxidine, lofexidine) is useful for the treatment of sialorrhea, administered by the paralingual, sublingual or buccal route. The patient to be treated is also given an antimuscarinic agent (e.g. oxybutynin, glycopyrrolate, ipratropium).

IT 286930-02-7, Fesoterodine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

 $(\alpha 2\text{-receptor agonist-antimus carinic agent combination for treatment of sialorrhea)$ 

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

REFERENCE COUNT:

3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2007:705973 CAPLUS

DOCUMENT NUMBER:

147:125829

TITLE:

Pharmaceutical combination comprising a PED5 inhibitor and a muscarinic antagonist for the treatment of LUTS Mastrell, Carl Erik Johan; Suesserman, Michael Allen

INVENTOR(S):

PATENT ASSIGNEE(S):

Pfizer Products Inc., USA

SOURCE:

PCT Int. Appl., 32pp.

DOCUMENT TYPE:

CODEN: PIXXD2

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

|       | PAI | ENT I | NO.                             |                                 |                                 | KIN                             | D                               | DATE                                   | ·<br>                           |                          | APPL:                    | ICAT:                    | ION I                    | ١٥.                      |                          | D                        | ATE                      |                          |
|-------|-----|-------|---------------------------------|---------------------------------|---------------------------------|---------------------------------|---------------------------------|--|---------------------------------|--------------------------|--------------------------|--------------------------|--------------------------|--------------------------|--------------------------|--------------------------|--------------------------|--------------------------|
|       |     | 2007  |                                 |                                 |                                 |                                 |                                 | 2007<br>2007                           |                                 | Ī                        | WO 2                     | 006-                     | IB368                    | 33                       |                          | 2                        | 0061                     | 219                      |
|       | WO  | W:    | AE,<br>CN,<br>GE,<br>KP,<br>MN, | AG,<br>CO,<br>GH,<br>KR,<br>MW, | AL,<br>CR,<br>GM,<br>KZ,<br>MX, | AM,<br>CU,<br>GT,<br>LA,<br>MY, | AT,<br>CZ,<br>HN,<br>LC,<br>MZ, | AU,<br>DE,<br>HR,<br>LK,<br>NA,<br>SG, | AZ,<br>DK,<br>HU,<br>LR,<br>NG, | DM,<br>ID,<br>LS,<br>NI, | DZ,<br>IL,<br>LT,<br>NO, | EC,<br>IN,<br>LU,<br>NZ, | EE,<br>IS,<br>LV,<br>OM, | EG,<br>JP,<br>LY,<br>PG, | ES,<br>KE,<br>MA,<br>PH, | FI,<br>KG,<br>MD,<br>PL, | GB,<br>KM,<br>MG,<br>PT, | GD,<br>KN,<br>MK,<br>RO, |
|       |     | R₩:   | AT,<br>IS,<br>CF,<br>GM,        | BE,<br>IT,<br>CG,<br>KE,        | BG,<br>LT,<br>CI,<br>LS,        | CH,<br>LU,<br>CM,<br>MW,        | CY,<br>LV,<br>GA,<br>MZ,        | VC,<br>CZ,<br>MC,<br>GN,<br>NA,<br>TM, | DE,<br>NL,<br>GQ,<br>SD,        | DK,<br>PL,<br>GW,<br>SL, | EE,<br>PT,<br>ML,<br>SZ, | ES,<br>RO,<br>MR,<br>TZ, | SE,<br>NE,               | SI,<br>SN,               | SK,<br>TD,               | TR,<br>TG,               | BF,<br>BW,               | BJ,<br>GH,               |
|       |     | 2007  | 1692                            | 78                              |                                 |                                 |                                 |  |                                 | ,                        | JP 2                     | 006-                     |                          |                          |                          |                          | 0061                     |                          |
| PRIOR | TT. | ( APP | LN.                             | INFO                            | .:                              |                                 |                                 |  |                                 |                          |                          |                          | 75262<br>7577:           |                          |                          |                          | 0051:<br>0060            |                          |
| GI    |     |       |                                 |                                 |                                 |                                 |                                 |  |                                 |                          |                          |                          |                          |                          |                          |                          |                          |                          |

This invention relates to the combined use of a phosphodiesterase 5 (PDE5) AΒ inhibitor and a muscarinic antagonist in the treatment of lower urinary tract symptoms (LUTS), such as urgency, frequency, nocturia and urge incontinence. A method of treatment of LUTS comprises simultaneous, sep., or sequential administration of a PED5 inhibitor and a muscarinic antagonist to a patient in need of such treatment. Thus, a muscarinic antagonist, oxybutynin (3.18 mg/kg) produced a small increase in micturition pressure, whereas the PED5 inhibitor, 3-ethyl-5-[5-(4ethylpiperazin-1-ylsulfonyl)-2-n-propoxyphenyl]-1-(pyridin-2-yl)methyl-1,6dihydro-7H-pyrazolo[4,3-d]pyrimidin-7-one (I, 0.11 mg/kg and 0.32 mg/kg) produced a small reduction in micturition pressure in guinea pigs. The combination of oxybutynin (3.18 mg/kg) plus I (0.32 mg/kg) produced a greater reduction in micturition pressure than observed with I (0.32 mg/kg) alone. These data appear to imply a synergistic effect of oxybutynin and the higher dose of I tested on micturition pressure. Also, an immediate-release tablet containing fesoterodine (muscarinic antagonist) and 5-[2-ethoxy-5-(4-ethylpiperazine-1-sulfonyl)pyridin-3-yl]-3-ethyl-2-(2methoxyethyl)-2,6-dihydropyrazolo[4,3-d]pyrimidin-7-one (PED5 inhibitor) were prepared comprising (i) a core containing fesoterodine hydrogen fumarate 2.0 mg, 5-[2-ethoxy-5-(4-ethylpiperazine-1-sulfonyl)pyridin-3-yl]-3-ethyl-

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2-(2-methoxyethyl)-2,6-dihydropyrazolo[4,3-d]pyrimidin-7-one besylate 5.0 mg, microcryst. cellulose 53.4 mg, calcium hydrogen phosphate dihydrate 18.0 mg, sodium starch glycollate 6.0 mg, magnesium stearate 0.4 mg, and colloidal silica 0.2 mg, and (ii) a coating containing methylhydroxypropyl cellulose 1.5 mg, microcryst. cellulose 0.3 mg, stearic acid 0.6 mg, and titanium dioxide E 171 0.6 mg.

IT 286930-02-7, Fesoterodine 286930-03-8

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(compns. comprising PED5 inhibitor and muscarinic antagonist for treatment of lower urinary tract disorders)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 286930-03-8 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester, (2E)-2-butenedioate (1:1) (CA INDEX NAME)

CM 1

CRN 286930-02-7 CMF C26 H37 N O3

Absolute stereochemistry. Rotation (+).

CM 2

CRN 110-17-8 CMF C4 H4 O4 Double bond geometry as shown.

L7 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2006:630212 CAPLUS

DOCUMENT NUMBER: 145:110309

TITLE: Injectable sustained release microspheric preparation

of 3,3-diphenylpropylamine derivatives as muscarinic

receptor antagonists

INVENTOR(S):
Li, Youxin

PATENT ASSIGNEE(S): Peop. Rep. China

SOURCE: PCT Int. Appl., 36 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: Chinese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PAT      | CENT  | NO.  | ٠    |     | KIN | D   | DATE |       |     | APPL | ICAT: | ION I | . 00   |     | D    | ATE  |     |
|----------|-------|------|------|-----|-----|-----|------|-------|-----|------|-------|-------|--------|-----|------|------|-----|
| WO       | 2006  | 0665 | 09   |     | A1  | -   | 2006 | 0629  |     | WO 2 | 005-0 | CN22  | <br>77 |     | 2    | 0051 | 222 |
| •        | W:    | ΑE,  | AG,  | AL, | AM, | ΑT, | ΑU,  | AZ,   | BA, | BB,  | BG,   | BR,   | BW,    | BY, | BZ,  | CA,  | CH, |
|          |       | CN,  | co,  | CR, | CU, | CZ, | DE,  | DK,   | DM, | DZ,  | EC,   | ΕE,   | EG,    | ES, | FI,  | GB,  | GD, |
|          |       | GE,  | GH,  | GM, | HR, | HU, | ID,  | IL,   | IN, | IS,  | JP,   | KE,   | KG,    | KM, | KN,  | KP,  | KR, |
|          |       | KZ,  | LC,  | LK, | LR, | LS, | LT,  | LU,   | LV, | LY,  | MA,   | MD,   | MG,    | MK, | MN,  | MW,  | MX, |
|          |       | MZ,  | NA,  | NG, | NI, | NO, | NZ,  | OM,   | PG, | PH,  | PL,   | PT,   | .RO,   | RU, | .SC, | SD,  | SE, |
|          |       | SG,  | SK,  | SL, | SM, | SY, | ТJ,  | TM,   | TN, | TR,  | TT,   | ΤZ,   | UA,    | UG, | US,  | UZ,  | VC, |
| •        |       | VN,  | YU,  | ZA, | ZM, | ·ZW |      |       |     |      |       |       |        |     |      |      |     |
|          | RW:   | AT,  | BE,  | BG, | CH, | CY, | CZ,  | DE,   | DK, | EE,  | ES,   | FI,   | FR,    | GB, | GR,  | HU,  | ΙE, |
|          |       | IS,  | IT,  | LT, | LU, | LV, | MC,  | NL,   | PL, | PT,  | RO,   | SE,   | SI,    | SK, | TR,  | BF,  | ВJ, |
|          |       | CF,  | CG,  | CI, | CM, | GΑ, | GN,  | GQ,   | GW, | ML,  | MR,   | NE,   | SN,    | TD, | TG,  | BW,  | GH, |
|          |       | GM,  | ΚE,  | LS, | MW, | ΜZ, | NA,  | SD,   | SL, | SZ,  | TZ,   | UG,   | ZM,    | ZW, | AM,  | ΑZ,  | BY, |
|          |       | KG,  | ΚZ,  | MD, | RU, | ТJ, | TM   |       |     |      |       |       |        |     |      |      |     |
| CN       | 1795  | 845  |      |     | Α   |     | 2006 | 0705  |     | CN 2 | 004-  | 1010  | 1721   |     | 2    | 0041 | 223 |
| PRIORITY | Y APP | LN.  | INFO | .:  |     |     |      |       |     | CN 2 | 004-  | 1010  | 1721   | i   | A 2  | 0041 | 223 |
| OTHER SO | DURCE | (S): |      |     | MAR | PAT | 145: | 11030 | 9   |      |       |       |        |     |      |      |     |

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AB The invention relates to injectable sustained release microspheric preparation of 3,3-diphenylpropylamine, its preparing process and application. The said sustained release microspheric preparation consists of 3,3-diphenylpropylamine of formula I as follows, its optical enantiomers or racemates and one or more medicinal biodegradable high-mol. auxiliary material and other medicinal auxiliary material, wherein the definition of R1, R2 R3 R4 and

R5 sees the claims. The injectable sustained release microspheric preparation according to the invention is used for treatment or supplementary treatment of diseases related to the muscarinic receptor and unstable or overactive bladder such as urgency or stress urinary incontinence, urge incontinence, urgency or frequency, etc.

IT 286930-02-7 895137-80-1

RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (injectable sustained release microspheric preparation of

3,3-diphenylpropylamine derivs. as muscarinic receptor antagonists)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 895137-80-1 CAPLUS

CN Benzenemethanol, 4-(acetyloxy)-3-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]- (CA INDEX NAME)

Absolute stereochemistry.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2006:76147 CAPLUS

DOCUMENT NUMBER: 144:156740

TITLE: Combinations of statins with bronchodilators for

treatment of respiratory disorders

INVENTOR(S): Lindmark, Bertil; Thoren, Anders Ingemar

PATENT ASSIGNEE(S): AstraZeneca AB, Swed.; AstraZeneca UK Limited

SOURCE: PCT Int. Appl., 18 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

|     | PAT                         | CENT I                          | NO.  |      |     | KIN | D   | DATE |      |     | APPL | ICAT  | ION  | NO. |     | D.  | ATE  |     |
|-----|-----------------------------|---------------------------------|------|------|-----|-----|-----|------|------|-----|------|-------|------|-----|-----|-----|------|-----|
|     | WO                          | 2006                            | 0084 | 37   |     | A1  | -   | 2006 | 0126 |     | WO 2 | 005-  | GB24 | 13  |     | 2   | 0050 | 620 |
| ٠.  |                             | W:                              | ΑE,  | AG,  | AL, | AM, | AT, | ΑU,  | ΑZ,  | BA, | ВВ,  | BG,   | BR,  | BW, | BY, | ΒŻ, | CA,  | CH, |
|     |                             |                                 | CN,  | CO,  | CR, | CU, | CZ, | DE,  | DK,  | DM, | DZ,  | EC,   | EE,  | EG, | ES, | FI, | GB,  | GD, |
|     |                             |                                 | GE,  | GH,  | GM, | HR, | HU, | ID,  | IL,  | IN, | IS,  | JP,   | ΚE,  | KG, | KM, | ΚP, | KR,  | ΚZ, |
|     |                             |                                 | LC,  | LK,  | LR, | LS, | LT, | LU,  | LV,  | MA, | MD,  | MG,   | MK,  | MN, | MW, | MX, | MZ,  | NA, |
|     |                             |                                 | NG,  | NI,  | NO, | ΝZ, | OM, | PG,  | PH,  | PL, | PT,  | RO,   | RU,  | SC, | SD, | SE, | SG,  | SK, |
|     |                             |                                 |      |      |     |     |     | TN,  |      |     |      |       |      |     |     |     |      |     |
|     |                             |                                 | ZA,  | ZM,  | ZW  |     |     |      |      |     |      | -     | -    | -   |     |     |      |     |
|     |                             | RW:                             | AT,  | BE,  | BG, | CH, | CY, | CZ,  | DE,  | DK, | EE,  | ES,   | FI,  | FR, | GB, | GR, | HU,  | IE, |
|     |                             |                                 | IS,  | IT,  | LT, | LU, | MC, | NL,  | PL,  | PT, | RO,  | SE,   | SI,  | SK, | TR, | BF, | ВJ,  | CF, |
|     |                             |                                 | CG,  | CI,  | CM, | GA, | GN, | GQ,  | GW,  | ML, | MR,  | NE,   | SN,  | TD, | TG, | BW, | GH,  | GM, |
|     |                             |                                 | KE,  | LS,  | MW, | MZ, | NA, | SD,  | SL,  | SZ, | ΤZ,  | UG,   | ZM,  | ZW, | AM, | ΑZ, | BY,  | KG, |
| ·   |                             |                                 | ΚZ,  | MD,  | RU, | ТJ, | TM  |      |      |     |      |       |      |     |     |     |      |     |
|     | KZ, MD, RU<br>AU 2005263883 |                                 |      |      |     | A1  |     | 2006 | 0126 |     | AU 2 | 005-  | 2638 | 83  |     | 2   | 0050 | 620 |
|     |                             | 2573                            |      |      |     |     |     |      |      |     |      |       |      |     |     |     | 0050 | 620 |
|     | ΕP                          | 1773                            | 319  |      |     | A1  |     | 2007 | 0418 |     | EP 2 | 005-  | 7520 | 46  |     | 2   | 0050 | 620 |
|     |                             | R:                              |      |      |     |     |     | CZ,  |      |     |      |       |      |     |     |     |      |     |
|     |                             |                                 | IS,  | IT,  | LI, | LT, | LU, | MC,  | NL,  | PL, | PT,  | RO,   | SE,  | SI, | SK, | TR, | AL,  | BA, |
|     |                             |                                 | HR,  | LV,  | MK, | ΥU  |     |      |      |     |      |       |      |     |     |     |      |     |
|     | CN 1984653                  |                                 |      |      |     |     |     | 2007 |      |     |      | 005-  |      |     |     | _   | 0050 |     |
|     | US 2008004247               |                                 |      |      |     |     |     | 2008 |      |     |      | 007-  |      |     |     |     | 0070 |     |
|     | MX 200700424                |                                 |      |      |     | Α   |     | 2007 |      |     |      | 007-  |      |     |     |     | 0070 |     |
|     |                             | KR 2007031392                   |      |      |     |     |     | 2007 |      | ,   | KR 2 | 007-  | 7008 | 31  |     | 2   | 0070 |     |
|     |                             | NO 2007000651<br>IN 2007DN01182 |      |      |     |     |     |      |      |     |      | 007-  |      |     |     | 2   | 0070 |     |
|     |                             |                                 |      |      |     | Α   |     | 2007 | 0427 |     |      | 007-1 |      |     |     |     | 0070 |     |
| RIO | RITY                        | APP.                            | LN.  | INFO | . : |     |     |      |      |     |      | 004-  |      |     |     |     | 0040 |     |
|     |                             |                                 |      |      |     |     |     |      |      | 1   | WO 2 | 005-0 | GB24 | 13  | 7   | W 2 | 0050 | 620 |

AB The invention provides medicaments comprising combinations of bronchodilators, glucocorticosteroids and HMG-CoA reductase inhibitors in the treatment of respiratory disorders such as chronic obstructive pulmonary disease (COPD). For example, a metered dose inhaler contained per dose formoterol fumarate dihydrate 4.5  $\mu g$ , budesonide 160  $\mu g$ , rosuvastatin 1 mg, and HFA 227 50  $\mu L$ . Also, an inhalation/oral combination comprised an aerosol formulation containing per dose formoterol fumarate dihydrate 4.5  $\mu g$  and budesonide 160  $\mu g$ , and a tablet formulation containing rosuvastatin 10 mg.

IT 286930-02-7, Fesoterodine

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combinations of statins with bronchodilators for treatment of respiratory disorders)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

CAPLUS COPYRIGHT 2008 ACS on STN L7ANSWER 9 OF 14

ACCESSION NUMBER:

2004:902168 CAPLUS

DOCUMENT NUMBER:

141:374727

TITLE:

Method using quaternary ammonium compounds for the

treatment of irritable bowel syndrome

INVENTOR(S):

Richards, Ivan Michael; Kolbasa, Karen Patrice

Pharmacia & Upjohn Company, LLC, USA

SOURCE:

PCT Int. Appl., 37 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION: DAMENIM NO

PATENT ASSIGNEE(S):

|                | PATENT NO. |              |            |            |            | KIN        | D          | DATE                     |            |            | APPL         | ICAT       | ION :      | NO.        |            | D.         | ATE          |            |
|----------------|------------|--------------|------------|------------|------------|------------|------------|--------------------------|------------|------------|--------------|------------|------------|------------|------------|------------|--------------|------------|
| *              |            | 2004<br>2004 |            |            |            | A2<br>A3   |            | 2004                     |            |            | WO 2         | 004-       | IB12       | 18         |            | 2          | 0040         | 405        |
|                |            | W:           | CN,<br>GE, | CO,<br>GH, | CR,<br>GM, | CU,<br>HR, | CZ,<br>HU, | AU,<br>DE,<br>ID,        | DK,<br>IL, | DM,<br>IN, | DZ,<br>IS,   | EC,<br>JP, | EE,<br>KE, | EG,<br>KG, | ES,<br>KP, | FI,<br>KR, | GB,<br>KZ,   | GD,<br>LC, |
|                |            | DW.          | NO,<br>TJ, | NZ,<br>TM, | OM,<br>TN, | PG,<br>TR, | PH,<br>TT, | LV,<br>PL,<br>TZ,<br>MW, | PT,<br>UA, | RO,<br>UG, | RU,<br>US,   | SC,<br>UZ, | SD,<br>VC, | SE,<br>VN, | SG,<br>YU, | SK,<br>ZA, | SL,<br>ZM,   | SY,<br>ZW  |
|                |            | KW:          | BY,<br>ES, | KG,<br>FI, | KZ,<br>FR, | MD,<br>GB, | RU,<br>GR, | TJ,<br>HU,<br>CG,        | TM,<br>IE, | AT,<br>IT, | BE,<br>LU,   | BG,<br>MC, | CH,<br>NL, | CY,<br>PL, | CZ,<br>PT, | DE,<br>RO, | DK,<br>SE,   | EE,<br>SI, |
| PRION<br>OTHER | RITY       |              | LN.        | 24<br>INFO |            | A1         |            | 2004                     |            |            | US 2<br>US 2 |            |            |            |            |            | 0040<br>0030 |            |
| GI             |            | 701101       | (0).       |            |            | 111111     |            | T 1 T                    | 0,11,      | <b>.</b> ' |              |            |            |            |            |            |              |            |

AΒ The invention discloses a method for treating irritable bowel syndrome by administering quaternary ammonium compds. Compds. of the invention include e.g. I [R1 = (un)] substituted C1-6 alkyl, (un) substituted CH2(C1-4 alkenyl), (un) substituted CH2(C1-6 alkynyl);  $\tilde{X}$  = anion of pharmaceutically acceptable acid]. Preparation of selected compds., e.g. (3R)-3-(2-hydroxy-5methylphenyl)-N, N-diisopropyl-N-methyl-3-phenylpropan-1-aminium bromide, is included.

Ι

ΙT 518360-93-5

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(quaternary ammonium compds. for treatment of irritable bowel syndrome)

518360-93-5 CAPLUS RN

CN Benzenepropanaminium, 5-(hydroxymethyl)-N-methyl-N, N-bis(1-methylethyl)-2- $(2-\text{methyl-1-oxopropoxy})-\gamma-\text{phenyl-}$ , bromide,  $(\gamma R)-(9CI)$  (CA) INDEX NAME)

## Absolute stereochemistry.

● Br-

L7 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2004:878361 CAPLUS

DOCUMENT NUMBER:

141:370546

TITLE:

Highly pure bases of 3,3-diphenyl propylamine monoesters for use in transdermal delivery

systems

INVENTOR(S):

Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael;

Drews, Roland

CODEN: PIXXD2

PATENT ASSIGNEE(S):

Schwarz Pharma Ag, Germany

PCT Int. Appl., 72 pp.

DOCUMENT TYPE:

LANGUAGE:

SOURCE:

Patent German

FAMILY ACC. NUM. COUNT:

: 1

PATENT INFORMATION:

| PAT | ENT          | NO.  |     | ٠   | KIN | D : | DATE  |      | •   | APPL | ICAT | ION I | NO.  |     | Di  | ATE  |     |    |
|-----|--------------|------|-----|-----|-----|-----|-------|------|-----|------|------|-------|------|-----|-----|------|-----|----|
| WO  | 2004         | 0898 | 72  |     | A1  |     | 2004  | 1021 | 1   | WO 2 | 004- | EP35  | 67   |     | 20  | 0040 | 403 |    |
|     | W:           | ΑE,  | AG, | AL, | AM, | AT, | AU,   | ΑZ,  | BA, | BB,  | BG,  | BR,   | BW,  | BY, | BZ, | CA,  | CH, |    |
|     |              | CN,  | CO, | CR, | CU, | CZ, | DE,   | DK,  | DM, | DZ,  | EC,  | EE,   | EG,  | ES, | FI, | GB,  | GD, |    |
|     |              | GE,  | GH, | GM, | HR, | HU, | ID,   | IL,  | IN, | IS,  | JP,  | ΚE,   | KG,  | KP, | KR, | ΚZ,  | LC, |    |
|     |              | LK,  | LR, | LS, | LT, | LU, | LV,   | MA,  | MD, | MG,  | MK,  | MN,   | MW,  | MX, | ΜŻ, | ΝA,  | NI, |    |
|     |              | NO,  | NΖ, | OM, | PG, | PH, | PL,   | PT,  | RO, | RU,  | SC,  | SD,   | SE,  | SG, | SK, | SL,  | SY, |    |
|     |              | ТJ,  | TM, | TN, | TR, | TT, | ΤZ,   | UA,  | UG, | US,  | UZ,  | VC,   | VN,  | YU, | ZA, | ZM,  | ZW  |    |
|     | RW:          | BW,  | GH, | GM, | KΕ, | LS, | MW,   | MΖ,  | SD, | SL,  | SZ,  | ΤZ,   | UG,  | ZM, | ZW, | ΑM,  | ΑZ, |    |
|     |              |      |     |     |     |     |       |      |     |      |      | CH,   |      |     |     |      |     |    |
|     |              |      |     |     |     |     |       |      |     |      |      | NL,   |      |     |     |      |     |    |
| •   |              |      |     |     |     | CF, | CG,   | CI,  | CM, | GA,  | GN,  | GQ,   | GW,  | ML, | MR, | ΝE,  | SN, |    |
|     |              |      |     |     |     |     |       |      |     |      |      |       |      |     |     |      |     |    |
|     | 1031         |      |     |     |     |     |       |      |     |      |      | 1031  |      |     |     | 0030 |     |    |
|     | 2004         |      |     |     |     |     | 2004  |      | 4   | AU 2 | 004- | 2281  | 63   |     | 2   | 0040 | 403 |    |
|     | 2004         |      |     |     |     |     | 2007  |      |     |      |      | 0505  |      |     | -   |      |     |    |
|     | 2505         |      |     |     | A1  |     |       |      |     |      |      | 2505  |      |     |     | 0040 |     |    |
|     | 2004         |      |     |     |     |     |       |      |     |      |      | 6221  |      |     | . 2 | 0040 |     |    |
|     | 1613         |      |     |     | A1  |     |       |      |     | EP 2 | 004- | 7256  | 10   |     | 2   | 0040 | 403 |    |
| EΡ  | 1613         |      |     |     |     |     |       |      |     |      |      |       |      |     |     |      |     |    |
|     | R:           | AT,  | BE, | CH, | DE, | DK, | ES,   | FR,  | GB, | GR,  | 1T,  | LI,   | LU,  | NL, | SE, | MC,  | PT, |    |
|     |              | IE,  | S1, | LT, | ĽV, | ŀΙ, | RO,   | MK,  | CY, | AL,  | TR,  | BG,   | CZ,  | EE, | HU, | PL,  | SK, | HR |
| CN  | 1802         | 345  | - 0 |     | A   |     | 2006  | 0/12 | ,   | CN 2 | 004- | 8000  | 9224 |     | 2   | 0040 | 403 |    |
| JP  | 2006<br>2005 | 5227 | 58  |     | T   |     | 2006. | 1005 |     | JP 2 | 006- | 5049  | 89   |     | 2   | 0040 | 403 |    |
| ZΑ  | 2005         | 0026 | 19  |     | А   |     | 2006  | 0426 |     | ZA 2 | 005- | 26/9  |      |     | 2   | 0050 | 331 |    |

| MX 2005PA03562         | Α  | 20050603 | MX | 2005-PA3562   |   | 20050401 |
|------------------------|----|----------|----|---------------|---|----------|
| US 2006014832          | A1 | 20060119 | US | 2005-532836   |   | 20050426 |
| NO 2005005078          | Α  | 20051031 | NO | 2005-5078     |   | 20051031 |
| PRIORITY APPLN. INFO.: |    |          | DE | 2003-10315917 | Α | 20030408 |
|                        |    |          | WO | 2004-EP3567   | W | 20040403 |

OTHER SOURCE(S):

GΙ

MARPAT 141:370546

AΒ The invention relates to a compound of general formula (I) wherein A represents deuterium or hydrogen, R represents a group selected from C1-6 alkyl, C3-10 cycloalkyl or Ph, which can be substituted by C1-3 alkoxy, fluorine, chlorine, bromine, iodine, nitro, amino, hydroxyl, oxo, mercapto or deuterium. The C atom marked with a \* (star) can be present in an (R) configuration, in an (S)-configuration or a mixture thereof. The invention is characterized in that the above-mentioned compds. are free bases with a degree of purity of more than 97 wt %. The invention also relates to a method for the production of highly pure compds. of general formula (I) and to the use thereof in the production of medicaments. Thus (R)-2-[3-(Diisopropylamino)-1-phenylpropyl}-4-(hydroxymethyl)phenol was reacted with isobutyric acid chloride to form fesoterodine. Fesoterodine was purified via the formation of its fumaric acid salt. 1.5 G of the highly pure fesoterodine was mixed with 8.5 g silicone adhesive Bio-PSA 7-4300 and applied to a foil in order to prepare a transdermal delivery system.

IT 286930-02-7P, Fesoterodine

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

IT 777075-72-6P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

RN 777075-72-6 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester, carbonate (1:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 286930-02-7 CMF C26 H37 N O3

Absolute stereochemistry. Rotation (+).

CM 2

CRN 463-79-6 CMF C H2 O3

REFERENCE COUNT:

THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN ACCESSION NUMBER: 2004:878163 CAPLUS

DOCUMENT NUMBER: 141:360690

Combination therapies of asthma, COPD, allergic and TITLE:

infectious rhinitis

INVENTOR(S): Richards, Ivan Michael; Manning, Robert Everett

PATENT ASSIGNEE(S): Pfizer Inc, USA

SOURCE: U.S. Pat. Appl. Publ., 20 pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PA      | TENT                               | NO.  |      |     | KIN |     | DATE |      |     | APPI | ICAT  | ION I | NO. |     | D<br>- | ATE  |     |
|---------|------------------------------------|------|------|-----|-----|-----|------|------|-----|------|-------|-------|-----|-----|--------|------|-----|
| US      | 2004                               | 2099 | 16   |     |     |     | 2004 | 1021 |     | US 2 | 2004- | 8243  | 15  |     | 2      | 0040 | 413 |
|         | 2522                               |      |      |     |     |     | 2004 | 1028 |     |      |       |       |     |     |        | 0040 | 405 |
| WO      | 2004                               |      |      |     |     |     |      |      |     |      |       |       |     |     |        | 0040 |     |
|         | 2004                               |      |      |     |     |     |      |      |     |      |       |       |     |     |        |      |     |
|         | W:                                 | ΑE,  | AG,  | AL, | AM, | AT, | AU,  | ΑZ,  | BA, | BB,  | BG,   | BR,   | BW, | BY, | BZ,    | CA,  | CH, |
|         |                                    | CN,  | CO,  | CR, | CÜ, | CZ, | DE,  | DK,  | DM, | DZ,  | EC,   | EE,   | EG, | ES, | FI,    | GB,  | GD, |
|         |                                    | GE,  | GH,  | GM, | HR, | HU, | ID,  | IL,  | IN, | IS,  | JP,   | KΕ,   | KG, | ΚP, | KR,    | ΚZ,  | LC, |
|         |                                    | LK,  | LR,  | LS, | LT, | LU, | LV,  | MA,  | MD, | MG,  | MK,   | MN,   | MW, | MX, | MZ,    | NA,  | NΙ, |
|         | NO, NZ, C<br>TJ, TM, T             |      |      |     | PG, | PH, | PL,  | PT,  | RO, | RU,  | SC,   | SD,   | SE, | SG, | SK,    | SL,  | SY, |
|         |                                    |      |      |     | •   |     |      | •    |     |      |       |       |     | •   | •      | -    |     |
|         | RW:                                |      |      |     |     |     | MW,  |      |     |      |       |       |     |     |        |      |     |
|         |                                    |      |      |     |     |     | ТJ,  |      |     |      |       |       |     |     |        |      |     |
| •       |                                    |      |      |     |     |     | HU,  |      |     |      |       |       |     |     |        |      |     |
|         |                                    |      |      | BF, | ВJ, | CF, | CG,  | CI,  | CM, | GΑ,  | GN,   | GQ,   | GW, | ML, | MR,    | NE,  | SN, |
|         |                                    | TD,  |      |     |     |     |      |      |     |      |       |       |     |     | _      |      |     |
| EP      | 1620                               |      |      |     |     |     |      |      |     |      |       |       |     |     |        |      |     |
|         | R:                                 |      |      |     |     |     | ES,  |      |     |      |       |       |     | ΝL, | SE,    | MC,  | PT, |
|         |                                    |      |      |     |     |     | TR,  |      |     |      |       |       |     |     | _      |      | 405 |
|         | BR 2004009492                      |      |      |     |     |     |      |      |     |      |       |       |     |     |        |      |     |
|         | JP 2006523674<br>MX 2005PA11225    |      |      |     |     |     |      |      |     |      |       |       |     |     |        | 0040 |     |
| MX      | MX 2005PA11225 ORITY APPLN. INFO.: |      |      |     |     |     | 2005 | 1214 |     |      |       |       |     |     |        | 0051 |     |
| PRIORIT | Y APP                              | LN.  | INFO | .:  |     |     |      |      |     |      | 2003- |       |     |     |        | 0030 |     |
|         |                                    |      |      |     |     |     |      |      |     | WO 2 | 2004- | IB11  | 70  | 1   | ₩ 2    | 0040 | 405 |
| OTHER S | OURCE                              | (S): | •    |     | MAR | PAT | 141: | 3606 | 90  |      |       |       |     |     |        |      |     |

The invention is directed to methods of treating asthma, COPD, allergic rhinitis, and infectious rhinitis by administering a first pharmaceutical agent including one or more compds. selected from the quaternary ammonium compds. (Markush structures are included) and a second pharmaceutical agent including one or more pharmaceutical agents selected from Adenosine A2a Receptor Agonists, D2-Dopamine Receptor Agonists, Phosphodiesterase Inhibitors (PDE's), corticosteroids, norepinephrine reuptake inhibitors, 4-hydroxy-7-[2-[2-[3-[2-phenylethoxy]-propylsulfonyl]ethylamino]ethyl]-1,3benzothiazol-2(3H)-one, and pharmaceutically acceptable salts thereof, and non-quaternized antimuscarinic compds.

IT 518360-93-5

> RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(combination therapies of asthma, COPD, allergic and infectious rhinitis)

518360-93-5 CAPLUS RN

Benzenepropanaminium, 5-(hydroxymethyl)-N-methyl-N, N-bis(1-methylethyl)-2-CN  $(2-\text{methyl-1-oxopropoxy})-\gamma-\text{phenyl-}$ , bromide,  $(\gamma R)-(9CI)$  (CA) INDEX NAME)

Absolute stereochemistry.

● Br-

L7 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:872676 CAPLUS

DOCUMENT NUMBER:

141:337790

TITLE:

Transdermal administration of (R)-3,3-

diphenylpropylamine monoesters

INVENTOR(S):

Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael;

Drews, Roland

PATENT ASSIGNEE(S):

SOURCE:

Schwarz Pharma Ag, Germany

PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

LANGUAGE:

Patent

German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PA | ГЕНТ         | NO.         |                   |     | KIN      | D . | DATE  |              | ;    | APPL | ICAT | ION          | NO.  |     | D   | ATE  |     |    |
|----|--------------|-------------|-------------------|-----|----------|-----|-------|--------------|------|------|------|--------------|------|-----|-----|------|-----|----|
| WO | 2004         | 0893        | <del></del><br>46 |     | A1       | _   | 2004: | 1021         | 1    | WO 2 | 004- | EP35         | 74   |     | 21  | 0040 | 403 |    |
|    | W:           | AE,         | AG,               | AL, | AM,      | AT, | ΑU,   | AZ,          | BA,  | BB,  | BG,  | BR,          | BW,  | BY, | ΒZ, | CA,  | CH, |    |
|    |              |             | ,                 | •   | •        | ,   | •     | DK,          |      | •    | •    |              | •    |     |     |      |     |    |
|    |              |             |                   |     |          |     |       | IL,          |      |      |      |              |      |     |     |      |     |    |
|    |              |             |                   |     | -        |     |       | MA,          |      |      |      |              |      |     |     |      |     |    |
|    |              |             |                   |     |          |     |       | PT,          |      |      |      |              |      |     |     |      |     |    |
|    | DEI.         |             |                   |     |          |     |       | UA,          |      |      |      |              |      |     |     |      |     |    |
|    | RW:          | BW,         | •                 | •   | •        |     |       | •            |      |      | •    | •            |      |     | •   |      |     |    |
|    |              |             | -                 |     |          |     |       | TM,<br>IE,   |      |      |      |              |      |     |     |      |     |    |
|    |              |             |                   |     |          | •   | •     | CI,          |      | •    |      |              |      | •   | •   |      | •   |    |
|    |              | TD,         | -                 | D., | Б0,      | 01, | 00,   | 01,          | 01.7 | J,   | 01., | 027          | ;    | ,   | ,   | ,    | J., |    |
| DE | 1031         | ,           |                   |     | A1       |     | 2004  | 1104         |      | DE 2 | 003- | 1031         | 5878 |     | 2   | 0030 | 408 |    |
| ΑU | 2004         | 2289        |                   |     |          |     |       |              |      |      |      |              |      |     |     | 0040 | 403 |    |
| ΑU | 2004         | 2289        | 27                |     | В2       |     | 2007  | 0517         |      |      |      |              |      |     |     |      |     |    |
| CA | 2505         | 780         |                   |     |          |     |       | 1021         |      |      |      | 2505         |      |     |     | 040  |     |    |
|    | 1530         |             |                   |     |          |     |       | 0518         |      | EP 2 | 004- | 7256         | 14   |     | 20  | 0040 | 403 |    |
| ΕP | 1530         |             |                   |     |          |     | 2007  |              |      |      |      |              |      |     |     |      |     |    |
|    | R:           | AT,         |                   |     |          |     |       |              |      |      |      |              |      |     |     |      |     |    |
|    | 0004         |             |                   |     |          |     |       | MK,          |      |      |      |              |      |     |     |      |     | нк |
|    | 2004         |             |                   |     |          |     | 2005  | 0816<br>0503 |      | CN 2 | 004- | 8000<br>8000 | 0176 |     | 21  | 2040 | 403 |    |
| UN | 1767<br>2006 | 02U<br>5227 | 50                |     | A.<br>Tr |     |       | 1005         |      |      |      |              |      |     |     |      |     |    |
| N7 | 5392         | 11          | 33                |     | Σ/<br>T  |     |       | 0223         |      |      |      |              |      |     |     | 0040 |     |    |
|    | 3746         |             |                   |     |          |     |       | 1015         |      |      |      | 7256         |      |     |     |      |     |    |
|    | 2005         |             |                   |     |          |     |       |              |      |      |      |              |      |     |     |      |     |    |
|    | 2005         |             |                   |     | A        |     |       | 1013         |      |      |      | 2681         |      |     |     |      |     |    |

US 2006029673 Α1 20060209 US 2005-533683 20050426 NO 2005004644 20051010 20051010 Α NO 2005-4644 PRIORITY APPLN. INFO.: 20030408 DE 2003-10315878 Α WO 2004-EP3574 20040403

OTHER SOURCE(S): GI

MARPAT 141:337790

Me

Me

Ι

The invention relates to a device for transdermally administering a compound of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by \* (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5 weight/weight%

ozokerite or ceresin was heated to  $150\,^\circ\text{C}$  for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at  $150\,^\circ\text{C}$ ; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies.

IT 286930-02-7P, Fesoterodine

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP (Physical process); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2003:950829 CAPLUS

DOCUMENT NUMBER: 140:13084

TITLE: Combination of selected opioids with other active

substances for use in the therapy of urinary

incontinence

INVENTOR(S):
Christoph, Thomas

PATENT ASSIGNEE(S): Grunenthal G.m.b.H., Germany

SOURCE: PCT Int. Appl., 126 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. |        |      |         |     | KIND DATE |       |      | APPLICATION NO. |     |    |       |      |        |     | DATE |      |         |  |
|------------|--------|------|---------|-----|-----------|-------|------|-----------------|-----|----|-------|------|--------|-----|------|------|---------|--|
| WO         | 2003   | 0992 | <b></b> |     | A1        | _     | 2003 | 1204            |     | WO | 2003- | EP55 | <br>29 |     | 2    | 0030 | <br>527 |  |
|            |        |      |         |     |           |       |      |                 |     |    | , BG, |      |        |     |      |      |         |  |
|            |        |      |         |     |           |       |      |                 |     |    | , ES, |      |        |     |      |      |         |  |
|            |        | HR,  | HU,     | ID, | IL,       | IN,   | IS,  | JP,             | KE, | KG | , KP, | KR,  | KZ,    | LC, | LK,  | LR,  | LS,     |  |
|            |        |      |         |     |           |       |      |                 |     |    | , MX, |      |        |     |      |      |         |  |
|            |        | PT,  | RO,     | RU, | SC,       | SD,   | SE,  | SG,             | SK, | SL | , TJ, | TM,  | TN,    | TR, | TT,  | TZ,  | UA,     |  |
|            |        | UG,  | US,     | UZ, | VC,       | VN,   | YU,  | ZA,             | ZM, | zw |       |      |        |     |      |      |         |  |
|            | RW:    | GH,  | GM,     | ΚE, | LS,       | MW,   | MZ,  | SD,             | SL, | SZ | , TZ, | UG,  | ZM,    | ZW, | AM,  | AZ,  | BY,     |  |
|            |        |      |         |     |           |       |      |                 |     |    | , CH, |      |        |     |      |      |         |  |
|            |        |      |         |     |           |       |      |                 |     |    | , NL, |      |        |     |      |      |         |  |
|            |        | BF,  | ВJ,     | CF, | CG,       | CI,   | CM,  | GA,             | GN, | GQ | , GW, | ML,  | MR,    | NE, | SN,  | TD,  | TG      |  |
| DE         | 1022   | 4107 |         |     | A1        |       | 2003 | 1211            |     | DE | 2002- | 1022 | 4107   |     | 2    | 0020 | 529     |  |
| AU         | 2003   | 2407 | 17      |     | A1        |       | 2003 | 1212            |     | ΑU | 2003- | 2407 | 17     |     | 2    | 0030 | 527     |  |
| EP         | 1507   | 520  |         |     | A1        |       | 2005 | 0223            |     | EΡ | 2003- | 7301 | 20     |     | 2    | 0030 | 527     |  |
|            | R:     | AT,  | BE,     | CH, | DE,       | DK,   | ES,  | FR,             | GB, | GR | , IT, | LI,  | LU,    | NL, | SE,  | MC,  | PT,     |  |
|            |        | ΙE,  | SI,     | LT, | LV,       | FΙ,   | RO,  | MK,             | CY, | AL | , TR, | BG,  | CZ,    | EE, | HU,  | SK   |         |  |
| US         | 2005   | 1371 | 94      |     | A1        |       | 2005 | 0623            |     | US | 2004- | 9981 | 64     |     | 2    | 0041 | 129     |  |
| US         | 2006   | 1689 | 42      |     | A1        |       | 2006 | 0803            |     | US | 2005- | 5459 | 01     |     | 2    | 0050 | 817     |  |
| US         | 7246   | 486  |         |     | В2        |       | 2007 | 0724            |     |    |       |      |        |     |      |      |         |  |
| PRIORITY   | Y APP  | LN.  | INFO    | . : |           |       |      |                 |     | DE | 2002- | 1022 | 4107   |     | A 2  | 0020 | 529     |  |
| •          |        |      |         |     |           |       |      |                 |     | WO | 2003- | EP55 | 29     |     | W 2  | 0030 | 527     |  |
| OMITTED 04 | 217000 | (0)  |         |     |           | - 7 m | 1 40 | 1 2 0 0         |     |    |       |      |        |     |      |      |         |  |

OTHER SOURCE(S): MARPAT 140:13084

AB The invention discloses the use of a combination of opioids (e.g. tramadol) with other active substances for producing a drug for the treatment of urinary urgency or urinary incontinence. The invention also relates to corresponding medicaments and to a method for treating urinary urgency or urinary incontinence.

IT 286930-02-7, Fesoterodine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL

(Biological study); USES (Uses)

(opioid combination with other active substances for treatment of urinary incontinence)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

REFERENCE COUNT: 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

1999:736261 CAPLUS

DOCUMENT NUMBER:

SOURCE:

131:336818

TITLE:

Preparation of 3,3-diphenylpropylamines as

antimuscarinic agents.

INVENTOR(S):

Sparf, Bengt; Meese, Claus O. Schwarz Pharma AG, Germany

PATENT ASSIGNEE(S):

Eur. Pat. Appl., 27 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PATENT NO.      | KIND DATE       | APPLICATION NO.         | DATE        |
|-----------------|-----------------|-------------------------|-------------|
| EP 957073       | A1 19991117     | EP 1998-108608          | 19980512    |
|                 |                 | GB, GR, IT, LI, LU, NL, |             |
|                 | LV, FI, RO      |                         |             |
|                 |                 | CA 1999-2328920         |             |
| WO 9958478      | A1 19991118     | WO 1999-EP3212          | 19990511    |
| W: AE, AL, AM,  | AT, AU, AZ, BA, | BB, BG, BR, BY, CA, CH, | CN, CU, CZ, |
| DE, DK, EE,     | ES, FI, GB, GD, | GE, GH, GM, HR, HU, ID, | IL, IN, IS, |
| JP, KE, KG,     | KP, KR, KZ, LC, | LK, LR, LS, LT, LU, LV, | MD, MG, MK, |
| MN, MW, MX,     | NO, NZ, PL, PT, | RO, RU, SD, SE, SG, SI, | SK, SL, TJ, |
|                 | UA, UG, US, UZ, |                         |             |
| RW: GH, GM, KE, | LS, MW, SD, SL, | SZ, UG, ZW, AT, BE, CH, | CY, DE, DK, |
| ES, FI, FR,     | GB, GR, IE, IT, | LU, MC, NL, PT, SE, BF, | BJ, CF, CG, |
|                 | GN, GW, ML, MR, |                         |             |
| AU 9941412      | A 19991129      | AU 1999-41412           | 19990511    |
| AU 748057       | B2 20020530     |                         |             |
| BR 9910406      | A 20010109      | BR 1999-10406           | 19990511    |
| EP 1077912      | A1 20010228     | EP 1999-924929          | 19990511    |
| EP 1077912      | B1 20020703     |                         |             |
| R: AT, BE, CH,  | DE, DK, ES, FR, | GB, GR, IT, LI, LU, NL, | SE, MC, PT, |
| IE, SI, LT,     | LV, FI, RO      |                         |             |
| ни 2001000779   | A2 . 20010828   | ни 2001-779             | 19990511    |
| TR 200003319    | T2 20011221     | TR 2000-3319            | 19990511    |

| AT        | 2200   | 56    |      |     | Т            |      | 2002    | 0715  | A.            | Г  | 1999-9 | 9249 | 29   |     |    | 199  | 905 | 511 |   |
|-----------|--------|-------|------|-----|--------------|------|---------|-------|---------------|----|--------|------|------|-----|----|------|-----|-----|---|
| EP        | 1254   | 890   |      |     | A1           |      | 2002    | 1106  | E             | 2  | 2002-3 | 1348 | 1    |     |    | 199  | 905 | 511 |   |
|           | R:     | AT,   | BE,  | CH, | DE,          | DK,  | ES,     | FR,   | GB, G         | GR | R, IT, | LI,  | LU,  | NL, | SI | Ξ, Μ | iC, | PT  | , |
|           |        | IE,   | SI,  | LT, | LV,          | ΓI,  | RO,     | MK,   | CY, A         | ٩L | ,      |      |      |     |    |      |     |     |   |
| NZ        | 5074   | 87    |      |     | Α            |      | 2002    | 1126  | N             | Z  | 1999-5 | 5074 | 87   |     |    | 199  | 905 | 511 |   |
| PT        | 1077   | 912   |      |     | T            |      | 2002    | 1129  |               |    | 1999-9 |      |      |     |    | 199  | 905 | 511 |   |
|           | 2181   |       |      |     | Т3           |      | 2003    | 0216  | ES            | 3  | 1999-9 | 9249 | 29   |     |    | 199  | 905 | 511 |   |
|           | 2199   |       |      |     | C2           |      | 2003    |       | RU            | J  | 2000-3 | 1258 | 13   |     |    | 199  |     |     |   |
| JP        | 2003   | 5190  | 79   |     | $\mathbf{T}$ |      | 2003    | 0617  | JI            | 2  | 2000-5 | 5482 | 84   |     |    | 199  | 905 | 511 |   |
| . JP      | 3929   | 702   |      |     | В2           |      | 2007    | 0613  |               |    |        |      |      |     |    |      |     |     |   |
| CN        | 1690   | 041   |      |     | Α            |      | 2005    | 1102  | Cì            | V  | 2005-3 | 1007 | 0299 |     |    | 199  | 905 | 511 |   |
| CZ        | 2966   | 05    |      |     | В6           |      | 2006    | 0412  | C             | Z  | 2000-3 | 3774 |      |     |    | 199  | 905 | 11  |   |
|           | 1955   |       |      |     | В1           |      | 2007    | 1031  |               |    | 1999-3 |      |      |     |    | 199  | 905 | 511 |   |
| SK        | 2860   | 52    |      |     | • в6         |      | 2008    | 0205  | SI            | <  | 2000-3 | 1547 |      |     |    | 199  | 905 | 511 |   |
| ZA        | 2000   | 0057  | 28   |     | · A          |      | 2001    | 0305  | $\mathbf{Z}I$ | Ą  | 2000-5 | 5728 |      |     |    | 200  | 010 | 17  |   |
| NO        | 2000   | 0056  | 69   |     | Α            |      | 2001    | 0111  | NO            | )  | 2000-5 | 5669 |      |     |    | 200  | 011 | 110 |   |
| MX        | 2000   | PA11  | 096  |     | Α            |      | 2002    | 0604  | MΣ            | ζ  | 2000-1 | PA11 | 096  |     |    | 200  | 011 | 110 |   |
| US        | 6713   | 464   |      |     | В1           | ٠    | 2004    | 0330  | US            | 3  | 2001-  | 7000 | 94   |     |    | 200  |     |     |   |
| HK        | 1046   | 269   |      |     | A1           |      | 2005    | 0923  | H             | <  | 2002-3 | 1078 | 59   |     |    | 200  |     |     |   |
| US        | 2004   | 1860  | 61   |     | A1           |      | 2004    | 0923  | US            | 3  | 2004-  | 7662 | 63   |     |    | 200  | 401 | L27 |   |
| US        | 7230   | 030   |      |     | В2           |      | 2007    | 0612  |               |    |        |      |      |     |    |      |     |     |   |
| US        | 2006   | 2707: | 38   |     | A1           |      | 2006    |       |               | _  | 2005-2 |      |      |     |    | 200  |     | -   |   |
|           | 2007   |       |      |     | Α            |      | 2007    |       |               |    | 2006-2 |      |      |     |    | 200  |     |     |   |
|           | 2007   |       |      |     | Α            |      | 2007    | 0816  |               | -  | 2007-3 |      |      |     |    | 200  |     |     |   |
| PRIORITY  | APP:   | LN.   | INFO | .:  |              |      |         |       |               | _  | 1998-  |      |      |     | Α  | 199  |     |     |   |
|           |        |       |      |     |              |      |         |       |               |    | 1999-8 |      |      |     |    | 199  |     |     |   |
|           |        |       |      |     |              |      |         |       |               |    | 1999-9 |      |      |     |    | 199  |     |     |   |
|           |        |       |      |     |              |      |         |       |               | _  | 2000-5 |      | -    |     |    | 199  |     |     |   |
|           |        |       |      |     |              |      |         |       |               | -  | 1999-I |      |      |     | W  | 199  |     |     |   |
|           |        |       |      |     |              |      |         |       |               |    | 2001-  |      |      |     |    | 200  |     |     |   |
|           |        |       |      |     |              |      |         |       |               | 5  | 2004-  | 7662 | 63   |     | A1 | 200  | 401 | L27 |   |
| OMILED CO | TIDOE. | 101.  |      |     | MADD         | עם ע | 7 7 7 . | 2260. | 10 .          |    |        |      |      |     |    |      |     |     |   |

OTHER SOURCE(S):

MARPAT 131:336818

Ι

AΒ Title compds. (I; R = H, Me, Et, Pr, Me2CH, Bu, iso-Bu, pentyl, hexyl, PhCH2, alkyl, CHO, Ac, propionyl, isobutyryl, aminocarbonyl, aminosulfonyl, MeO2C, etc.; R1 = H, Me, Et, Pr, Me2CH, Bu, iso-Bu, pentyl, hexyl, PhCH2, alkyl, phenylalkyl; Z = NR8R9; R8, R9 = hydrocarbyl; NR8R9 = atoms to form a ring; with a proviso), were prepared as antimuscarinic agents (no data). Thus, 4-bromophenol, cinnamoyl chloride, and Et3N were stirred 18 h in CH2Cl2 to give 99.8% 3-phenylacrylic acid 4-bromophenyl ester. This was refluxed 2 h with HOAc/H2SO4 to give 43.8% 6-bromo-4-phenylchroman-2-one. The latter was refluxed with benzyl bromide, K2CO3, and NaI in acetone/MeOH to give 102.1% crude Me 3-(2-benzyloxy-5-bromophenyl)-3-phenylpropionate, which was stirred with LiAlH4 in THF to give 96.3% 3-(2-benzyloxy-5-bromophenyl)-3-phenylpropan-1ol. This was stirred with tosyl chloride and pyridine in CH2Cl2 for 18 h to give 93.6% tosylate ester, which was refluxed 97 h with diisopropylamine in MeCN to give 77.9% [3-(2-benzyloxy-5-bromophenyl)-3phenylpropyl]diisopropylamine. The latter was converted in several steps to 2-(3-diisopropylamino-1-phenylpropyl)-4-hydroxymethylphenol, which was acylated to give I.

RN 250214-42-7 CAPLUS
CN Benzenemethanol, 3-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(1-oxopropoxy)- (CA INDEX NAME)

RN 250214-43-8 CAPLUS
CN Butanoic acid, 2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4(hydroxymethyl)phenyl ester (CA INDEX NAME)

RN 250214-44-9 CAPLUS
CN Propanoic acid, 2-methyl-, 2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

RN 250214-45-0 CAPLUS

CN Propanoic acid, 2,2-dimethyl-, 2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

RN 250214-46-1 CAPLUS

CN Benzenemethanol, 4-(benzoyloxy)-3-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]- (CA INDEX NAME)

RN 250214-47-2 CAPLUS

CN Propanedioic acid, bis[2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl] ester (9CI) (CA INDEX NAME)

RN 250214-48-3 CAPLUS

CN Butanedioic acid, bis[2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl] ester (9CI) (CA INDEX NAME)

RN 250214-49-4 CAPLUS

Pentanedioic acid, bis[2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-CN (hydroxymethyl)phenyl] ester (9CI) (CA INDEX NAME)

RN 250214-50-7 CAPLUS

Hexanedioic acid, bis[2-[3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-CN (hydroxymethyl)phenyl] ester (9CI) (CA INDEX NAME)

REFERENCE COUNT:

3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

# => D L6 IBIB ABS HITSTR 1-3

ANSWER 1 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2007:1454781 CAPLUS

DOCUMENT NUMBER:

148:78876

TITLE:

SOURCE:

Cyclopentylpyrrolidinone derivatives and their preparation and use in combination therapy for the treatment of urinary frequency, urinary urgency and

urinary incontinence

INVENTOR(S):

Gottesdiener, Keith M.; Green, Stuart A.; Macintyre,

Euan

PATENT ASSIGNEE(S):

Merck & Co., Inc., USA PCT Int. Appl., 86pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT      | PATENT NO.    |     |      |     | KIND DATE |     |      |      | APPL | ICAT |      | DATE |     |     |      |      |     |
|----------|---------------|-----|------|-----|-----------|-----|------|------|------|------|------|------|-----|-----|------|------|-----|
| WO       | WO 2007146224 |     |      |     |           | _   | 2007 | 1221 | 1    | WO 2 | 007- | US13 | 683 |     | 21   | 0070 | 607 |
|          | W:            | ΑE, | AG,  | AL, | AM,       | AT, | ΑU,  | ΑZ,  | BA,  | BB,  | BG,  | BH,  | BR, | BW, | BY,  | ΒZ,  | CA, |
|          |               | CH, | CN,  | CO, | CR,       | CU, | CZ,  | DE,  | DK,  | DM,  | DO,  | DZ,  | EC, | EE, | EG,  | ES,  | FI, |
|          |               | GB, | GD,  | GE, | GH,       | GM, | GT,  | HN,  | HR,  | HU,  | ID,  | IL,  | IN, | IS, | JP,  | ΚE,  | KG, |
|          |               | KM, | KN,  | KP, | KR,       | ΚZ, | LA,  | LC,  | LK,  | LR,  | LS,  | LT,  | LU, | LY, | MA,  | MD,  | ME, |
|          |               | MG, | MK,  | MN, | MW,       | MX, | MY,  | ΜZ,  | NA,  | NG,  | NI,  | NO,  | NΖ, | OM, | PG,  | PH,  | PL, |
|          |               | PT, | RO,  | RS, | RU,       | SC, | SD,  | SE,  | SG,  | SK,  | SL,  | SM,  | SV, | SY, | ТJ,  | TM,  | TN, |
|          |               | TR, | TT;  | ΤZ, | UA,       | UG, | US,  | UZ,  | VC,  | VN,  | ZA,  | ZM,  | ZW  |     |      |      |     |
|          | RW:           | ΑT, | BE,  | BG, | CH,       | CY, | CZ,  | DE,  | DK,  | EE,  | ES,  | FI,  | FR, | GB, | GR,  | ΗU,  | ΙE, |
|          |               | IS, | IT,  | LT, | LU,       | LV, | MC,  | MT,  | NL,  | PL,  | PT,  | RO,  | SE, | SI, | SK,  | TR,  | BF, |
|          |               | ВJ, | CF,  | CG, | CI,       | CM, | GΑ,  | GN,  | GQ,  | GW,  | ML,  | MR,  | NE, | SN, | TD,  | ΤG,  | B₩, |
|          |               | GH, | GM,  | ΚE, | LS,       | MW, | MZ,  | NA,  | SD,  | SL,  | SZ,  | TZ,  | UG, | ZM, | ZW,  | AM,  | ΑZ, |
|          |               | BY, | KG,  | ΚZ, | MD,       | RU, | ТJ,  | TM   |      |      |      |      |     |     |      | _    |     |
| PRIORITY | APP:          | LN. | INFO | . : |           |     |      |      |      | US 2 | 006- | 8127 | 43P | 1   | P 20 | 0060 | 612 |
| GT       |               |     |      |     |           |     |      |      |      |      |      |      |     |     |      |      |     |

This invention concerns compns. for the treatment of urinary frequency, urinary urgency and urinary incontinence comprising a selected antagonist of the NK-1 receptor or a pharmaceutically acceptable salt thereof and an anti-muscarinic agent or a pharmaceutically acceptable salt thereof. This invention concerns combination therapy for urinary frequency, urinary urgency and urinary incontinence wherein one of the active agents is a selected antagonist of the NK-1 receptor or a pharmaceutically acceptable salt thereof and another is an anti-muscarinic agent or a pharmaceutically acceptable salt thereof. Example compound I was prepared by a multistep procedure (procedure given). All the invention compds. were evaluated for their NK-1 receptor antagonistic activity.

IT 286930-02-7, Fesoterodine

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(preparation of cyclopentylpyrrolidinone derivs. as anti-muscarinic agents and NK-1 receptor antagonists in combination therapy of urinary frequency, urinary urgency and urinary incontinence)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

L6 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:878361 CAPLUS

DOCUMENT NUMBER: 141:370546

TITLE: Highly pure bases of 3,3-diphenyl propylamine

monoesters for use in transdermal delivery

systems

INVENTOR(S): Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael;

Drews, Roland

PATENT ASSIGNEE(S): Schwarz Pharma Ag, Germany

SOURCE: PCT Int. Appl., 72 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: German

FAMILY ACC. NUM. COUNT: PATENT INFORMATION:

GΙ

|          | PATENT NO.   |      |        |     |          |     |      |      | APPLICATION NO. |       |       |          |        |     | DATE |       |       |    |
|----------|--------------|------|--------|-----|----------|-----|------|------|-----------------|-------|-------|----------|--------|-----|------|-------|-------|----|
|          | 2004         |      | <br>72 | •   | A1       | -   | 2004 | 1021 |                 | WO 2  | 2004- | <br>EP35 | <br>67 |     | 2    | 0040  | 403   |    |
|          | W:           | ΑE,  | AG,    | AL, | AM,      | AT, | AU,  | ΑZ,  | BA,             | BB,   | BG,   | BR,      | BW,    | ΒY, | BZ,  | CA,   | CH,   |    |
|          |              | CN,  | CO,    | CR, | CU,      | CZ, | DE,  | DK,  | DM,             | DZ,   | EC,   | EE,      | EG,    | ES, | FI,  | GB,   | GD,   |    |
|          |              |      |        |     |          |     | ID,  |      |                 |       |       |          |        |     |      |       |       |    |
|          |              |      |        |     |          |     | LV,  |      |                 |       |       |          |        |     |      |       |       |    |
|          |              |      |        | -   |          |     | PL,  | -    | -               | -     |       |          |        |     |      |       |       |    |
|          |              |      | -      |     |          |     | TZ,  | -    |                 |       |       | -        | -      | -   |      |       |       |    |
|          | RW:          |      |        | •   |          | •   | MW,  |      |                 |       | •     |          |        |     |      |       |       |    |
|          |              |      |        |     |          |     | TJ,  |      |                 |       |       |          |        |     |      |       |       |    |
|          |              |      |        |     |          |     | HU,  |      |                 |       |       |          |        |     |      |       |       |    |
|          |              |      |        | Br, | ΒJ,      | CF, | CG,  | CI,  | CM,             | GA,   | GN,   | GQ,      | GW,    | ML, | MK,  | NE,   | SN,   |    |
| DE       | 1031         | TD,  |        |     | 7.1      |     | 2004 | 1110 |                 | מם    | 0003  | 1021     | 5017   |     | 2    | 0020  | 4 N O |    |
|          | 2004         |      |        |     | A1       |     |      |      |                 |       |       |          |        |     |      |       |       |    |
|          | 2004         |      |        |     |          |     |      |      |                 | AU Z  | 2004- | 2201     | 0,5    |     | 2    | 0040  | 403   |    |
|          | 2505         | 848  | 05     |     | D2<br>Δ1 |     | 2007 | 1021 |                 | C A 2 | 004-  | 2505     | 848    |     | 2    | 0040  | 403   |    |
|          | 2004         | 0062 | 21     | ,   | Δ        |     | 2005 | 0809 |                 | BR 2  | 2004  | 6221     | 040    |     | 2    | 0040  | 403   |    |
|          | 1613         |      |        |     |          |     |      |      |                 |       |       |          |        |     |      |       |       |    |
| EP       | 1613         | 584  |        |     | B1       |     | 2007 | 1121 |                 |       |       | , 200    |        |     | _    |       |       |    |
|          |              |      |        |     |          |     | ES,  |      |                 | GR.   | IT.   | LI.      | LU,    | NL. | SE.  | ·MC,  | PT,   |    |
|          |              | IE,  | SI,    | LT, | LV,      | FI, | RO,  | MK,  | CY,             | AL,   | TR,   | BG,      | CZ,    | EE, | HU,  | PL,   | SK,   | HR |
| CN       | 1802         | 345  | •      | ·   | A.       | •   | 2006 | 0712 | ·               | CN 2  | 2004- | 8000     | 9224   |     | 2    | 0040  | 403   |    |
| JP       | 1802<br>2006 | 5227 | 58     |     | T        |     | 2006 | 1005 |                 | JP 2  | 2006- | 5049     | 89     |     | 2    | 0040  | 403   |    |
| ZA       | 2005         | 0026 | 79     |     | Α        |     | 2006 | 0426 |                 | ZA 2  | 2005- | 2679     |        |     | 2    | 0050. | 331   |    |
| MX       | 2005         | PA03 | 562    |     | Α        |     | 2005 | 0603 |                 | MX 2  | 2005- | PA35     | 62     |     | 2    | 0050  | 401   |    |
| US       | 2006         | 0148 | 32     |     | Al       |     | 2006 | 0119 |                 | US 2  | 2005- | 5328     | 36     |     | 2    | 0050  | 426   |    |
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|          |              |      |        |     |          |     |      |      |                 | WO 2  | 2004- | EP35     | 67     |     | W 2  | 0040  | 403   |    |
| OTHER SO | OURCE        | (S): |        |     | MAR      | PAT | 141: | 3705 | 46              |       |       |          |        |     |      |       |       |    |

AΒ The invention relates to a compound of general formula (I) wherein A represents deuterium or hydrogen, R represents a group selected from C1-6 alkyl, C3-10 cycloalkyl or Ph, which can be substituted by C1-3 alkoxy, fluorine, chlorine, bromine, iodine, nitro, amino, hydroxyl, oxo, mercapto or deuterium. The C atom marked with a \* (star) can be present in an (R) configuration, in an (S)-configuration or a mixture thereof. The invention is characterized in that the above-mentioned compds. are free bases with a degree of purity of more than 97 wt %. The invention also relates to a method for the production of highly pure compds. of general formula (I) and to the use thereof in the production of medicaments. Thus (R)-2-[3-(Diisopropylamino)-1-phenylpropyl]-4-(hydroxymethyl)phenol was reacted with isobutyric acid chloride to form fesoterodine. Fesoterodine was purified via the formation of its fumaric acid salt. 1.5 G of the highly pure fesoterodine was mixed with 8.5 g silicone adhesive Bio-PSA 7-4300 and applied to a foil in order to prepare a transdermal delivery system.

IT 286930-02-7P, Fesoterodine

RL: PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

IT 777075-72-6P

RL: PRP (Properties); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(highly pure bases of 3,3-di-Ph propylamine monoesters for use in transdermal delivery systems)

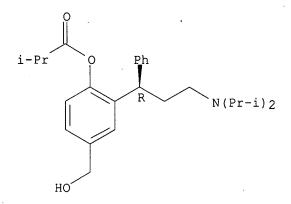
RN 777075-72-6 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester, carbonate (1:1) (salt) (9CI) (CA INDEX NAME)

CM 1

CRN 286930-02-7 CMF C26 H37 N O3

Absolute stereochemistry. Rotation (+).



CM 2

CRN 463-79-6 CMF C H2 O3

О || НО— С— ОН

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2004:872676 CAPLUS

DOCUMENT NUMBER:

141:337790

TITLE:

Transdermal administration of

(R)-3, 3-diphenylpropylamine monoesters

INVENTOR(S):

Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael;

Drews, Roland

PATENT ASSIGNEE(S):

Schwarz Pharma Ag, Germany

SOURCE:

PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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| WO 2004 | 0893 | 46  |     | A1  | •   | 2004 | 1021 | Ţ   | WO 2 | 004- | EP35  | 74  |     | 2   | 0040 | 403 |
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PRIORITY APPLN. INFO.:
                                                   DE 2003-10315878
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                                                                          W
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OTHER SOURCE(S): GΙ

MARPAT 141:337790

The invention relates to a device for transdermally administering a compound AΒ of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by \* (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of formula (I) for producing transdermal medicaments. silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5 weight/weight% ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies. ΙT

286930-02-7P, Fesoterodine

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP (Physical process); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

RN 286930-02-7 CAPLUS

> Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS-RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

#### => D L5 IBIB ABS HITSTR 1

ANSWER 1 OF 1 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2004:872676 CAPLUS

DOCUMENT NUMBER:

141:337790

TITLE:

Transdermal administration of (R)-3,3-

diphenylpropylamine monoesters

INVENTOR(S):

Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael;

Drews, Roland

PATENT ASSIGNEE(S):

Schwarz Pharma Ag, Germany

SOURCE:

PCT Int. Appl., 68 pp.

DOCUMENT TYPE:

CODEN: PIXXD2

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

| PAT                                      | PATENT NO. |       |     |     | KIND DATE |     |       | APPLICATION NO. |     |      |       |      |      | DATE |      |       |     |  |
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| AT       | 374605        | T       | 20071015    | AT 2004-725614      |     | 20040403       |
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OTHER SOURCE(S):

MARPAT 141:337790

GΙ

AB The invention relates to a device for transdermally administering a compound of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by  $\star$  (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through human skin. The invention further relates to the use of said compds. of formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5 weight/weight% ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies.

IT 286930-02-7P, Fesoterodine

RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP (Physical process); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses) (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

RN 286930-02-7 CAPLUS

CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1-phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

Ι

REFERENCE COUNT:

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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Executing the logoff script...

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100.0% PROCESSED

84 ITERATIONS

19 ANSWERS

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COST IN U.S. DOLLARS

SINCE FILE ENTRY

TOTAL SESSION

FULL ESTIMATED COST

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31 L2 L3

=> S L3 AND SKIN

274112 SKIN

1 L3 AND SKIN

=> D L4 IBIB ABS HITSTR 1

ANSWER 1 OF 1 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER:

2004:872676 CAPLUS.

DOCUMENT NUMBER:

141:337790

TITLE:

Transdermal administration of (R)-3,3-

diphenylpropylamine monoesters

INVENTOR(S):

Breitenbach, Armin; Meese, Claus; Wolff, Hans-Michael;

Drews, Roland

PATENT ASSIGNEE(S):

Schwarz Pharma Ag, Germany

SOURCE:

PCT Int. Appl., 68 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

### PATENT INFORMATION:

| PATENT NO.   |   | KIND  | DATE  | APPLICATION NO.   | DATE  |
|--|---|---|---|---|---|
| W: AE, AG, CN, CO, GE, GH, LK, LR, NO, NZ, TJ, TM, RW: BW, GH, BY, KG, ES, FI, SK, TR, | AL,<br>CR,<br>GM,<br>LS,<br>OM,<br>TN,<br>GM,<br>KZ,<br>FR, | AM, AT, CU, CZ, HR, HU, LT, LU, PG, PH, TR, TT, KE, LS, MD, RU, GB, GR, | AU, AZ, DE, DK, ID, IL, LV, MA, PL, PT, TZ, UA, MW, MZ, TJ, TM, HU, IE, | WO 2004-EP3574 BA, BB, BG, BR, BW, BY, DM, DZ, EC, EE, EG, ES, IN, IS, JP, KE, KG, KI, MD, MG, MK, MN, MW, MY, RO, RU, SC, SD, SE, SC, UG, US, UZ, VC, VN, YC, SD, SL, SZ, TZ, UG, ZN, AT, BE, BG, CH, CY, CZ, IT, LU, MC, NL, PL, PC, CM, GA, GN, GQ, GW, MI | Y, BZ, CA, CH, S, FI, GB, GD, P, KR, KZ, LC, K, MZ, NA, NI, G, SK, SL, SY, J, ZA, ZM, ZW A, ZW, AM, AZ, Z, DE, DK, EE, F, RO, SE, SI, |
| TD, TG DE 10315878 AU 2004228927 AU 2004228927   |   | A1  |   | DE 2003-10315878<br>AU 2004-228927  | 20030408<br>20040403  |
| CA 2505780<br>EP 1530461   |   | A1  | 20041021  | CA 2004-2505780<br>EP 2004-725614   |   |
| R: AT, BE,<br>IE, SI,<br>BR 2004006212   | CH,<br>LT,  | DE, DK,<br>LV, FI,<br>A   | RO, MK,<br>20050816   | GB, GR, IT, LI, LU, NI<br>CY, AL, TR, BG, CZ, ER<br>BR 2004-6212<br>CN 2004-80009176  | E, HU, PL, SK, HR<br>20040403   |
| JP 2006522759<br>NZ 539214<br>AT 374605  |   | T<br>A<br>T   | 20061005<br>20070223<br>20071015  | JP 2006-504992<br>NZ 2004-539214<br>AT 2004-725614  | 20040403<br>20040403<br>20040403  |
| ZA 2005002681<br>US 2006029673<br>NO 2005004644  |   | A<br>A1   | 20051013<br>20060209  | MX 2005-PA3561<br>ZA 2005-2681<br>US 2005-533683<br>NO 2005-4644<br>DE 2003-10315878  | 20050401<br>20050426<br>20051010  |
| PRIORITY APPLN. INFO.  OTHER SOURCE(S):  |   | MARPAT  | 141:3377  | WO 2004-EP3574  |   |

GΙ

The invention relates to a device for transdermally administering a compound of formula (I), wherein A represents hydrogen or deuterium, R represents a group selected among C1-6 alkyl, C3-10 cycloalkyl, or Ph, each of which can be substituted by C1-3 alkoxy, fluoride, chlorine, bromine, iodine, nitro, amino, hydroxy, oxo, mercapto, or deuterium, the C atom marked by \* (asterisk) being provided in the R configuration. The invention is characterized in that the compound of general formula (I) is provided in a polymer matrix and is released at a dose of 0.5 to 20 mg per day through

human skin. The invention further relates to the use of said compds. of formula (I) for producing transdermal medicaments. Thus a silicone-based transdermal system was prepared by the hot-melt process. 8.5 G of an adhesive mixture composed of BIO-PSA 7-4300 from Dow-Corning and 5 weight/weight% ozokerite or ceresin was heated to 150°C for 20 min until a homogeneous melt was formed. 1.5 G fesoterodine were added to the melt; the mixture was kept for addnl. 5 min at 150°C; followed by application onto a preheated foil. 5 Cm2 samples were used for dissoln. studies.

IT 286930-02-7P, Fesoterodine
 RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP
 (Physical process); SPN (Synthetic preparation); THU (Therapeutic use);
 BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
 (transdermal administration of (R)-3,3-diphenylpropylamine monoesters)

RN 286930-02-7 CAPLUS
CN Propanoic acid, 2-methyl-, 2-[(1R)-3-[bis(1-methylethyl)amino]-1phenylpropyl]-4-(hydroxymethyl)phenyl ester (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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SINCE FILE COST IN U.S. DOLLARS TOTAL ENTRY SESSION 188.23 FULL ESTIMATED COST 8.53 SINCE FILE TOTAL DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) ENTRY SESSION -0.80 CA SUBSCRIBER PRICE -0.80

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